

IN THE UNITED STATES PATENT AND TRADEMARK OFFICE

In re the application of: Curtis, Rory A.J.

Serial No.: 09/587,111

Filed: June 2, 2000

For: NOVEL MEMBERS OF THE  
CAPSAICIN/VANILLOID RECEPTOR FAMILY OF  
PROTEINS AND USES THEREOF

Attorney Docket No.: MNI-062CP2DV1



Group Art Unit: 1646

Examiner: Ulm, J. D.

TECH CENTER 1600/2900

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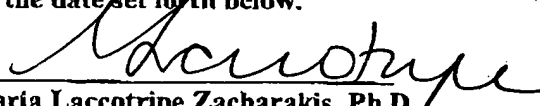
Certificate of Facsimile Transmission

I hereby certify that this correspondence is being facsimile transmitted to the Commissioner for Patents,  
Box AF, Washington, D.C. 20231 on the date set forth below.

4/11/03

Date of Signature

By:

  
Maria Laccotripe Zacharakis, Ph.D.  
Attorney for Applicant  
Limited Recognition Under 37 C.F.R. 10.9(b)

DECLARATION PURSUANT TO 37 CFR §1.131

Dear Sir:

I, Rory A.J. Curtis, a citizen of the United Kingdom, residing at 78 Hardwick Road,  
Ashland, Massachusetts 01721 hereby declare as follows:

(1) I am the inventor of the subject matter described and claimed in the above-identified application.

(2) Prior to January 22, 1999, the invention described and claimed in the above-referenced patent application was completed in this country, as evidenced by the following:

(a) Prior to January 22, 1999 I had completed the sequencing of the full length human VR-2 (SEQ ID NO:5) as evidenced by copies of the assembled contig map and the complete sequence of SEQ ID NO:5 derived from the contig map. The contig map and sequence print-out are submitted herewith as Exhibits A and B, respectively.

(b) Prior to January 22, 1999 I had determined that the polypeptide of SEQ ID NO:5 was a vanilloid receptor based on the results of a blast search using the amino acid sequence of SEQ ID NO:5. The top hit in this blast search analysis is rat vanilloid receptor 1 (VR-1; accession number AF029310). Based on the percent identity and percent similarity between the polypeptide of SEQ ID NO:5 and rat VR-1, and the fact that rat VR-1 was the top hit in the blast search analysis I determined that the polypeptide of SEQ ID NO:5 was a vanilloid receptor. The results of the blast search analysis are submitted herewith as Exhibit C.

(c) Prior to January 22, 1999, I had determined that the polypeptide of SEQ ID NO:5 represents a unique target for pain and that it may be responsible for hypersensitivity in chronic neuropathic pain, as evidenced by a copy of the "Qualified Target Summary Sheet" generated for this molecule. The Qualified Target Summary Sheet is submitted herewith as Exhibit D.

(d) According to my routine practice, immediately after identifying the polypeptide of SEQ ID NO:5 as a vanilloid receptor (hVR-2) associated with pain and classifying it as a Qualified Target, I contemplated routine uses of this molecule such as: (i) use of the molecule in screening assays to identify modulators of the vanilloid receptor or (ii) use of this molecule in the diagnosis of conditions/diseases associated with, for example, aberrant vanilloid receptor (hVR-2) nucleic acid expression or activity, *e.g.*, pain disorders, as evidenced by a copy of the "Qualified Target Summary Sheet" generated for this molecule.

(e) The dates deleted from the contig map (Exhibit A), the sequence print-out (Exhibit B), the blast search results (Exhibit C), and the Qualified Target Summary (Exhibit D) are prior to January 22, 1999.

I hereby declare that all statements made herein of my own knowledge are true and that all statements made on information and belief are believed to be true; and further that these statements are made with the knowledge that willful false statements and the like so made are punishable by fine or imprisonment, or both, under Section 1001 of Title 18 of the United States Code and that such willful false statements may jeopardize the validity of this Application for Patent or any patent issuing thereon.

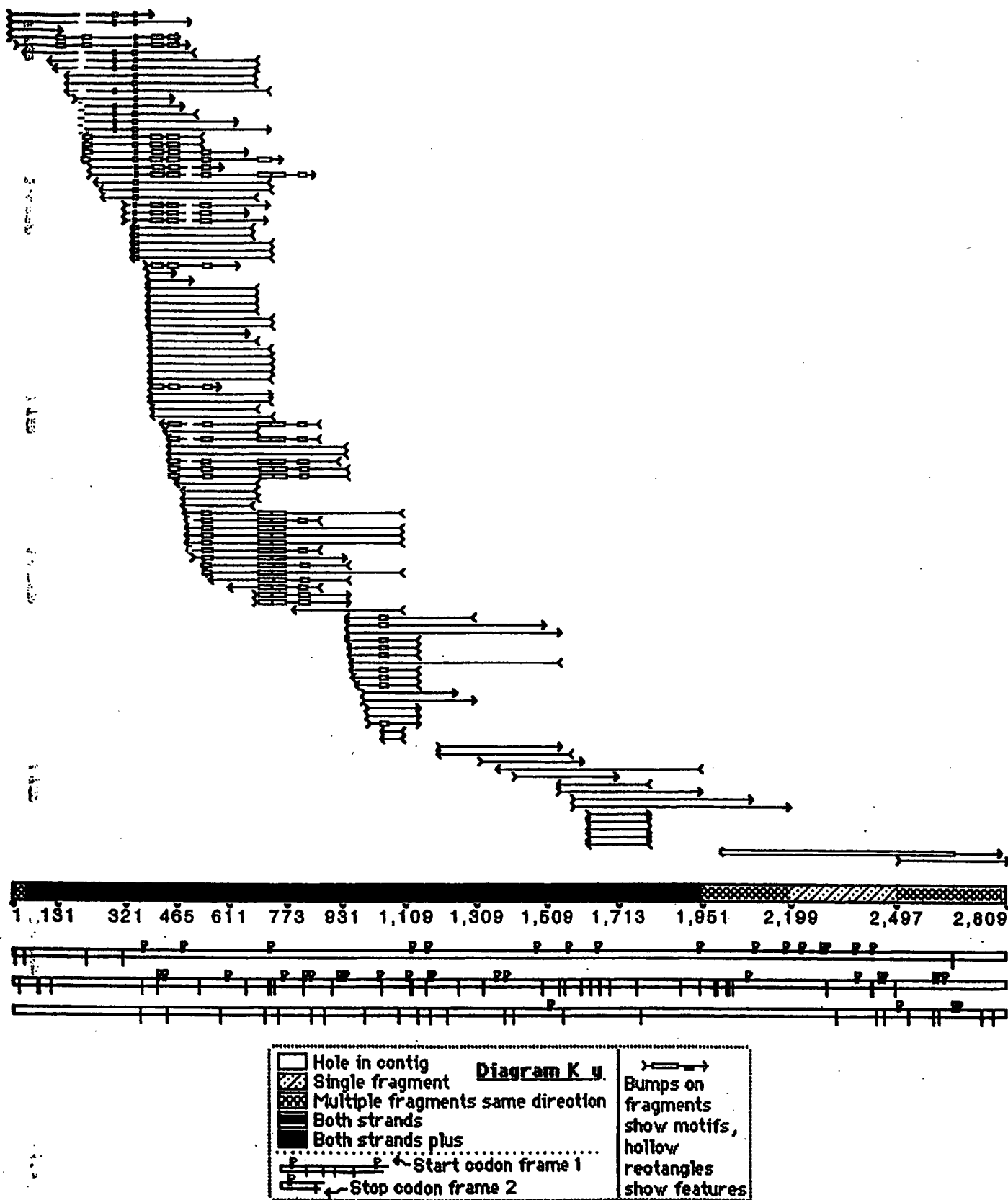
Rory Curtis  
Rory A.J. Curtis, Ph.D.

4/10/2003  
Date

**Exhibit A**

21e11a

Sequencher™ "21e11racefinal"



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21ella
Sequencher™ "21ellracefinal"

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\* \* + + + +

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+ . . . . + + + . . . + +

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+ + . + + + + + + + + + + + + + + + +

21ella  
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+ . . . . + + + + + + + + + + + + + + + + + +

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[illegible]

21ella  
Sequencher™ "21ellracefinal"

103e2w2 >#1>

**TG**

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\* + \* .. .... \* + \* + \* + \* + \* + \* + .. \* + \*



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Sequencher™ "21e11racefinal"

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flhX21e11r15g1	#67	GATTTTGGGA								

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jthsa103e2w1 #4	NGAAAGGGAA	CAGGTGCCAG	TCAGCCGGAT	CCAAACCGAT	TTGACCGAGA	TCGGCTCTTC	AATGCGGTCT	CCCGGGGTGT	CCCCGAGGAT
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#541 CGAAAGGGAA CAGGTGCCAG TCAGCCGGAT CCAAACCGAT TTGACCGAGA TCGGCTCTTC AATGCGGTCT CCCGGGGTGT CCCCGAGGAT  
R K G T G A S Q P D P N R F D R D R L F N A V S R G V P E D  
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flhX21e11r15..	#243	CTGGCTGGAC	TTCCAGAGTA	CCTGAGCAAG	ACCAGCAAAGT	ACCTCACCGA	CTCGGAATAC	ACAGAGGGCT	CCACAGGTAA	GACGTGCCTG
flhX21e11r15..	#242	CTGGCTGGAC	TTCCAGAGTA	CCTGAGCAAG	ACCAGCAAAGT	ACCTCACCGA	CTCGGAATAC	ACAGAGGGCT	CCACAGGTAA	GACGTGCCTG
flhX21e11r18..	#242	CTGGCTGGAC	TTCCAGAGTA	CCTGAGCAAG	ACCAGCAAAGT	ACCTCACCGA	CTCGGAATAC	ACAGAGGGCT	CCACAGGTAA	GACGTGCCTG
flhX21e11r16..	#241	CTGGCTGGAC	TTCCAGAGTA	CCTGAGCAAG	ACCAGCAAAGT	ACCTCACCGA	CTCGGAATAC	ACAGAGGGCT	CCACAGGTAA	GACGTGCCTG
flhX21e11r17..	#240	CTGGCTGGAC	TTCCAGAGTA	CCTGAGCAAG	ACCAGCAAAGT	ACCT: NCCGA	CTCGGAATAC	ACAGANGGCT	TCACANGTNA	AACGTGCCTT
flhX21e11r17..	#240	CTGGCTGGAC	TTCCAGAGTA	CCTGAGCAAG	ACCAGCAAAGT	ACCTCACCGA	CTCGGAATAC	ACAGAGGGCT	CCACAGGTAA	GACGTGCCTG
flhX21e11r16..	#239	CTGGCTGGAC	TTCCAGAGTA	CCTGAGCAAG	ACCAGCAAAGT					

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flhX1e11rr9c...	#89	CTGGCTGGAC	TTCCAGAGTA	CCTGAGCAAG	ACCAGCAAGT	ACCTCACCGA	CTCGGAATAC	ACAGAGGGCT	CCACAGGGCA	GACGTGCCTG
jthsa103e2x2	#74	CTGGCTGGAC	TTCCAGAGTA	CCTGAGCAAG	ACCAGCAAGT	ACCTCACCGA	CTCGGAATAC	ACAGAGGGCT	CCACAGGTAA	GACGTGCCTG
Genbank AI00...	#23	CTGGCTGGAC	TTCCAGAGTA	CCTGAGCAAG	ACCAGCAAGT	ACCTCACCGA	CTCGGAATAC	ACAGAGGGCT	CCACAGGTAA	GACGTGCCTG
jthsa103e2a1	>#1>						CTCGGAATAC	ACAGAGGGCT	CCACAGGTAA	GACGTGCCTG
jthsa103e2a2	>#1>						CTCGGAATAC	ACAGAGGGCT	CCACAGGTAA	GACGTGCCTG

#631 CTGGCTGGAC TTCCAGAGTA CCTGAGCAAG ACCAGCAAGT ACCTCACCGA CTCGGAATAC ACAGAGGGCT CCACAGGTAA GACGTGCCCTG  
 L A G L P E Y L S K T S K Y L T D S E Y T E G S T G K T C L  
 + + + + + + + + + + + + + + + + + + +

|                |      |  |
|----------------|------|--|
| F1hXc21e11h1   | #560 | AT:AAGGCTG TGCT  |
| F1hXc21e11b2   | #520 | ATGAAGGCTG TGCT  |
| jthsa103e2t1   | #511 | ATGAAGGCTG TGCTGAACCT TAAGGACNGA GTCAATGCCT GCATTNTGC  |
| jthsa103e02t.. | #500 | ATGAAGGCTG TGCTGAACCT TAAGGACNGA GTCAATGCCT GCATTCTGCC ACTGCTGCAG ATCGACAGGG ACTNTGQCAA TCCTNAGCCC |
| F1hX21e11r21.. | #485 | ATGAAGGCTG TGCTGAAC  |
| F1hX21e11r21.. | #466 | ATGAAGGCTG TGCTGAACC   |
| Genbank H20..  | #401 | ATGAAGGCTG TG  |
| Genbank H49..  | #400 | ATGAA  |
| F1hX21e11r22.. | #382 | ATGAAGGCTG TGCTGAACC   |
| F1hX21e11r22.. | #382 | ATGAAGGCTG TGCTGAACC   |
| F1hX21e11r20.. | #382 | ATGAAGGCTG TGCTGAACC   |
| F1hX21e11r20.. | #337 | ATGAAGGCTG TGCTGAACC   |
| F1hX21e11r18.. | #337 | ATGAAGGCTG AGCTGAACC   |
| F1hX21e11r17.. | #333 | ATGAAGGCTG TGCTGAAC  |
| F1hX21e11r15.. | #333 | ATGAAGGCTG TGCTGAACC   |
| F1hX21e11r15.. | #332 | ATGAAGGCTG TGCTGAACC   |
| F1hX21e11r18.. | #332 | ATGAAGGCTG AGCTGAACC   |
| F1hX21e11r16.. | #331 | ATGAAGGCTG TGCTGAAC  |
| F1hX21e11r17.. | #330 | ATGAANGCTT TGCTGAAC  |
| F1hX21e11r17.. | #330 | ATGAAGGCTG TGCTGAAC  |
| F1hX21e11r16.. | #329 | ATGAAGGCTG TGCTGAACC   |
| Genbank AA7..  | #302 | ATGAAGGCTG TGCTGAACCT TAAGGACGGA GTCAATGCCT GCATTCTGCC ACTGCTGCAG ATCGACAGGG ACTCTGGCAA TCCTCAGCCC |
| Genbank AA7..  | #279 | ATGAAGGCTG TGCTGAACCT TAAGGACGGG GTCAATGCCT GCATTCTGCC ACTGCTGCAG ATCGACAGGG ACTCTGGCAA TCCTCAGCCC |
| F1hX21e11r19.. | #278 | ATGAAGGCTG TGCTGAACCT TAAGGACGGG GTCAATGCCT GCATTCTGCC ACTGCTGCAG ATCGACAGGG ACTCTGGCAA TCCTCAGCCC |
| F1hX21e11r19.. | #278 | ATGAAGGCTG TGCTGAACCT TAAGGACGGG GTCAATGCCT GCATTCTGCC ACTGCTGCAG ATCGACAGGG ACTCTGGCAA TCCTCAGCCC |
| Genbank AIL..  | #276 | ATGAAGGCTG TGCTGAACCT TAAGGACGGA GTCAATGCCT GCATTCTGCC ACTGCTGCAG ATCGACAGGG ACTCTGGCAA TCCTCAGCCC |
| jthsa103e2w2   | #273 | ATGAAGGCTG TGCTGAACCT TAAGGACGGA GTCAATGCCT GCATTCTGCC ACTGCTGCAG ATCGACAGGG ACTCTGGCAA TCCTCAGCCC |
| jthsa103e2x1   | #268 | ATGAAGGCTG TGCTGAACCT TAAGGACGGA GTCAATGCCT GCATTCTGCC ACTGCTGCAG ATCGACAGGG ACTCTGGCAA TCCTCAGCCC |
| F1hX21e11rr1.. | #234 | ATGAAGGCTG TGCTGAACCT TAAGGACGGA GTCAATGCCT GCATTCTGCC ACTGCTGCAG ATCGACAGGG ACTCTGGCAA TCCTCAGCCC |
| Genbank AA4..  | #231 | ATGAAGGCTG TGCTGAACCT TAAGGACGGA GTCAATGCCT GCATTCTGCC ACTGCTGCAG ATCGACAGGG ACTCTGGCAA TCCTCAGCCC |
| F1hX21e11rr1.. | #231 | ATGAAGGCTG TGCTGAACCT TAAGGACGGA GTCAATGCCT GCATTCTGCC ACTGCTGCAG ATCGACAGGG ACTCTGGCAA TCCTCAGCCC |
| F1hX21e11rr1.. | #230 | ATGAAGGCTG TGCTGAACCT TAAGGACGGA GTCAATGCCT GCATTCTGCC ACTGCTGCAG ATCGACAGGG ACTCTGGCAA TCCTCAGCCC |
| F1hX21e11rr1.. | #229 | ATGAAGGCTG TGCTGAACCT TAAGGACGGG GTCAATGCCT GCATTCTGCC ACTGCTGCAG ATCGACAGGG ACTCTGGCAA TCCTCAGCCC |
| Genbank AA8..  | #224 | ATGAAGGCTG TGCTGAACCT TAAGGACGGG GTCAATGCCT GCATTCTGCC ACTGCTGCAG ATCGACAGGG ACTCTGGCAA TCCTCAGCCC |
| Genbank AA2..  | #213 | ATGAAGGCTG TGCTGAACCT TAAGGACGGG GTCAATGCCT GCATTCTGCC ACTGCTGCAG ATCGACAGGG ACTCTGGCAA TCCTCAGCCC |
| jthsa103e2w1   | #184 | ATGAAGGCTG TGCTGAACCT TAAGGACGGA GTCAATGCCT GCATTCTGCC ACTGCTGCAG ATCGACAGGG ACTCTGGCAA TCCTCAGCCC |
| F1hX21e11rr5.. | #179 | ATGAAGGCTG TGCTGAACCT TAAGGACGGA GTCAATGCCT GCATTCTGCC ACTGCTGCAG ATCGACAGGG ACTCTGGCAA TCCTCAGCCC |
| jthsa103e2x2   | #164 | ATGAAGGCTG TGCTGAACCT TAAGGACGGA GTCAATGCCT GCATTCTGCC ACTGCTGCAG ATCGACAGGG ACTCTGGCAA TCCTCAGCCC |
| Genbank AI0..  | #113 | ATGAAGGCTG TGCTGAACCT TAAGGACGGG GTCAATGCCT GCATTCTGCC ACTGCTGCAG ATCGACAGGG ACTCTGGCAA TCCTCAGCCC |
| jthsa103e2a1   | #41  | ATGAAGGCTG TGCTGAACCT TAAGGACGGA GTCAATGCCT GCATTCTGCC ACTGCTGCAG ATCGACAGGG ACTCTGGCAA TCCTCAGCCC |
| jthsa103e2a2   | #41  | ATGAAGGCTG TGCTGAACCT TAAGGACGGA GTCAATGCCT GCATTCTGCC ACTGCTGCAG ATCGACAGGG ACTCTGGCAA TCCTCAGCCC |
| F1hX21e11rr1.. | >1>  |  |

#721 ATGAAGGCTG TGCTGAACCT TAAGGACGGA GTCAATGCCT GCATTCTGCC ACTGCTGCAG ATCGACAGGG ACTCTGGCAA TCCTCAGCCC  
M K A V L N L K D G V N A C I L P L L Q I D R D S G N P Q P  
. + . . + . + . . + . . + .

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jthsa103e02t... #590 CTGGTAAATG CCCAGTGCAC AGATGACTAT TACCGAGGCC ACAGC C
Genbank AA7... #392 CTGGTAAATG CCCAGTGCAC AGATGACTAT TACCGAGGCC ACAGCGCTCT GCACATCGCC ATT
Genbank AA7... #369 CTGGTAAATG CCCAGTGCAC AGATGACTAT TACCGAGGCC ACAGCG:TCT GCACATCGCC ATT
FlhK21e11r19... #368 CTGGTAAATG CCCAGTGCAC AGATGACTAT TACCGAGGCC ACAGCGCTCT GCACATCGCC ATTGAGAAGA GGAGTCTGCA GTGTGTGAAG
FlhK21e11r19... #368 CTGGTAAATG CCCAGTGCAC AGATGACTAT TACCGAGGCC ACAGCGCTCT GCACATCGCC ATTGAGAAGA GGAGTCTGCA GTGTGTGAAG
Genbank AIL... #366 CTGGTAAATG CCCAGTGCAC AGATGACTAT TACCGAGGCC ACAGCGCTCT GCACATCGCC ATTGAGAAGA GGAGTCTGCA GTGTGTGAAG
jthsa103e2w2... #363 CTGGTAAATG CCCAGTGCAC AGATGACTAT TACCGAGGCC ACAGCGCTCT GCACATCGCC ATTGAGAAGA GGAGTCTGCA GTGTGTGAAG
jthsa103e2x1... #358 CTGGTAAATG CCCAGTGCAC AGATGACTAT TACCGAGGCC ACAGCGCTCT GCACATCGCC ATTGAGAAGA GGAGTCTGCA GTGTGTGAAG
FlhK21e11rr1... #324 CTGGTAAATG CCCAGTGCAC AGATGACTAT TACCGAGGCC ACAGCGCTCT GCACATCGTC ATTGAGAAGA GGAGTCTGCA GTGTGTGAAG
Genbank AA4... #321 CTGGTAAATG CCCAGTGCAC AGATGACTAT TACCGAGGCC ACAGCGCTCT GCACATCGCC ATT
FlhK21e11rr1... #321 CTGGTAAATG CCCAGTGCAC AGATGACTAT TACCGAGGCC ACAGCGCTCT GCACATCGCC ATTGAGAAGA GGAGTCTGCA GNGTGTNAAG
FlhK21e11rr1... #320 CTGGTAAATG CCCAGTGCAC AGATGACTAT TACCGAGGCC ACAGCGCTCT GCACATCGCC ATTGAGAAGA GGAGTCTGCA GTGTGTGAAG
FlhK21e11rr1... #319 CTGGTAAATG CCCAGTGCAC AGATGACTAT TACCGAGGCC ACAGCGCTCT GCACATCGCC ATTGAGAAGA GGAGTCTGCA GTGTGTGAAG
Genbank AA8... #314 CTGGTAAATG CCCAGTGCAC AGATGACTAT TACCGAGGCC ACAGCGCTCT GCACATCGCC ATT
Genbank AA2... #303 CTGGTAAATG CCCAGTGCAC AGATGACTAT TACCGAGGCC ACAGCGCTCT GCACATCGCC ATTGAGAAGA GGAGTCTGCA GTGTGTGAAG
jthsa103e2w1... #274 CTGGTAAATG CCCAGTGCAC AGATGACTAT TACCGAGGCC ACAGCGCTCT GCACATCGCC ATTGAGAAGA GGAGTCTGCA GTGTGTGAAG
FlhK21e11rr9... #269 CTGGTAAATG CCCAGTGCAC AGATGACTAT TACCGAGGCC ACAGCGCTCT GCACATCGTC ATTGAGAAGA GGAGTCTGCA GTGTGTGAAG
jthsa103e2x2... #254 CTGGTAAATG CCCAGTGCAC AGATGACTAT TACCGAGGCC ACAGCGCTCT GCACATCGCC ATTGAGAAGA GGAGTCTGCA GTGTGTGAAG
Genbank AIO... #203 CTGGTAAATG CCCAGTGCAC AGATGACTAT TACCGAGGCC ACAGCGCTCT GCACATCGCC ATT
jthsa103e2a1... #131 CTGGTAAATG CCCAGTGCAC AGATGACTAT TACCGAGGCC ACAGCGCTCT GCACATCGCC ATTGAGAAGA GGAGTCTGCA GTGTGTGAAG
jthsa103e2a2... #131 CTGGTAAATG CCCAGTGCAC AGATGACTAT TACCGAGGCC ACAGCGCTCT GCACATCGCC ATTGAGAAGA GGAGTCTGCA GTGTGTGAAG
FlhK21e11rr14... #25 CTGGTAAATG CCCAGTGCAC AGATGACTAT TACCGAGGCC ACAGCGCTCT GCACATCGCC ATTGAGAAGA GGAGTCTGCA GTGTGTGAAG

#811 CTGGTAAATG CCCAGTGCAC AGATGACTAT TACCGAGGCC ACAGCGCTCT GCACATCGCC ATTGAGAAGA GGAGTCTGCA GTGTGTGAAG
      L V N A Q C T D D Y Y R G H S A L H I A I E K R S L Q C V K
                + + . . + +

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#18 flhX21e11r19... #458 CTCCTGGTGG AGAATGGGGC CAATGTGCAT GCCCGGGCCT GCGGCCGC
#18 flhX21e11r19... #458 CTCCTGGTGG AGAATGGGGC CAATGTGCAT GCCCGGGCCT GCGGCCGC
#18 Genbank A11... #456 CTCCTGGTGG AGAATGGGGC CAATG
#18 jthsa103e2w2 #453 CTCCTGGTGG AGAATGGGGC CAATGTGCAT GCCCGGGCCT GCGGCCGCCT
#18 jthsa103e2w1 #448 CTCCTGGTGG AGAATGGGGC CAATGTGCAT GCCCGGGCCT GCGGCCGCCT
#18 flhX21e11rr1... #414 CTCCTGGTGG AGAATGGGGC CAATGTGCAT GCCCGGGCCT GCGACCGCTT CTTCCAGAAG GGCCAAGGGA CTTGCTTTTA TTTCGGTGAG
#18 flhX21e11rr1... #411 CTCCTGGTGG AGAATGGGGC CAATGTGCAT GCCCGGGCCT GCGGCCGCTT NTTCCAGAAG GGCCAAGGGA CTTGCTTTTA TTTCGGTNAG
#18 flhX21e11rr1... #410 CTCCTGGTGG AGAATGGGGC CAATGTGCAT GCCCGGGCCT GCGGCCGCTT CTTCCAGAAG GGCCAAGGGA CTTGCTTTTA TTTCGGTGAG
#18 flhX21e11rr1... #409 CTCCTGGTGG AGAATGGGGC CAATGTGCAT GCCCGGGCCT GCGGCCGCTT CTTCCAGAAG GGCCAAGGGA CTTGCTTTTA TTTCGGTGAG
#18 Genbank AA2... #393 CTCCTGGTGG AGAATGGGGC CAATGTGCAT GCCCGGGCCT G
#18 jthsa103e2w1 #364 CTCCTGGTGG AGAATGGGGC CAATGTGNAT GCCCGGGCCT GCGGCCGCCT
#18 flhX21e11rr9... #359 CTCCTGGTGG AGAATGGGGC CAATGTGCAT GCCCGGGCCT GCGACCGCTT CTTCCAGAAG GGCCAAGGGA CTTGCTTTTA TTTCGGTGAG
#18 jthsa103e2w2 #344 CTCCTGGTGG AGAATGGGGC CAATGTGCAT GCCCGGGCCT GCGGCCGCCT
#18 jthsa103e2a1 #221 CTCCTGGTGG AGAATGGGGC CAATGTGCAT GCCCGGGCCT GCGGCCGCCT
#18 jthsa103e2a2 #221 CTCCTGGTGG AGAATGGGGC CAATGTGCAT GCCCGGGCCT GCGGCCGCCT
#18 flhX21e11rr1... #115 CTCCTGGTGG AGAATGGGGC CAATGTGCAT GCCCGGGCCT GCGGCCGCTT CTTCCAGAAG GGCCAAGGGA CTTGCTTTTA TTTCGGTGAG
#18 frhob12c4g1... >#1> GCGGCCGCTT CTTCCAGAAG GGCCAAGGGA CTTGCTTTTA TTTCGGTGAG
#18 frhob012c04a... >#1> GCGGCCGCTT CTTCCAGAAG GGCCAAGGGA CTTGCTTTTA TTTCGGTGAG
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#18 flhX21e11rha2 >#1> CCGCCGCTT CTTCCAGAAG GGCCAAGGGA CTTGCTTTTA TTTCGGTGAG
#18 flhX21e11rmv2 >#1> GCGCCTT CTTCCAGAAG GGCCAAGGGA CTTGCTTTTA TTTCGGTGAG
#18 flhX21e11r1v1 >#1> CCGCCTT CTTCCAGAAG GGCCAAGGGA CTTGCTTTTA TTTCGGTGAG
#18 frhoc12c4h3... >#1> TT CTTCCAGAAG GGCCAAGGGA CTTGCTTTTA TTTCGGTNAG
#18 flhX21e11r1v2 >#1> TTCCAGAAG GGCCAAGGGA CTTGCTTTTA TTTCGGTGAG
#18 flhX21e11rmv1 >#1> AGAAG GGCCAAGGGA CTTGCTTTTA TTTCGGTGAG
#18 flhX21e11rja2 >#1> AGGGA CTTGCTTTTA TTTCGGTGAG
#18 Genbank AA9... >#1> TCGGTGAG
#18 Genbank AA9... >#1> TCGGTGAG

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#901 CTCCTGGTGG AGAATGGGGC CAATGTGCAT GCCCGGGCCT GCGGCCGCTT CTTCCAGAAG GGCCAAGGGA CTTGCTTTTA TTTCGGTGAG
      L L V E N G A N V H A R A C G R F F Q K G Q G T C F Y F G E
          + . + . + . + . +

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## 21e11a

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|                 |      |            |            |            |            |            |            |            |            |            |
|-----------------|------|------------|------------|------------|------------|------------|------------|------------|------------|------------|
| flhX21ellrr1..  | #504 | CTACCCCTCT | CTTTGGCCGC | TTGCACCAAG | CAGTGGGATG | TGGTAAGCTA | CCTCCTGGAG | AACCCACACC | AGCCCCCCAG | CCTGCAGGCC |
| flhX21ellrr1..  | #501 | CTACCCCTCT | CTTTGGCCGC | TTGCACCAAG | CAGTGGGATG | TGGTAAGCTA | CCTCCTGGAG | AACCCACACC | AGCCCNCCAG | CCTGCAGGCC |
| flhX21ellrr1..  | #500 | CTACCCCTCT | CNTTGGCCGC | TTGCACCAAG | CAGTGGGATG | TGGTAAGCTA | CCTCCTGGAG | AACCCACACC | AGCCCCCCAG | CCTGCAGGCC |
| flhX21ellrr1..  | #499 | CTACCCCTCT | CTTTGGCCGC | TTGCACCAAG | CAGTGGGATG | TGGTAAGCTA | CCTCCTGGAG | AACCCACACC | GGCCCCCCAG | CCTGCAGGCC |
| flhX21ellrr9..  | #449 | CTACCCCTCT | CTTTGGCCGC | TTGCACCAAG | CAGTGGGATG | TGGTAAGCTA | CCTCCTGGAG | AACCCACACC | AGCCCCCCAG | CCTGCAGGCC |
| flhX21ellrr1..  | #205 | CTACCCCTCT | CTTTGNCCGC | TTGCACCAAG | CAGTGGGATG | TGGTAAGCTA | CCTCCTGGAG | AACCCACACC | GGCCCCCCAG | CCTGCAGGCC |
| frhob12c4g1..   | #51  | CNACCCCTCT | CTTTGNCCGC | TTGCACCAAG | CAGTGGNATG | TGGTAAGCTA | CCTCCTGGAG | AACCCACNCC | AGCCCCCCAG | CCTGCAGGCC |
| frhob012c04s1.. | #51  | CTACCCCTCT | CTTTGGCCGC | TTGCACCAAG | CAGTGGGATG | TGGTAAGCTA | CCTCCTGGAG | AACCCACACC | AGCCCCCCAG | CCTGCAGGCC |
| frhob012c04s1.. | #51  | CTACCCCTCT | CTTTNGCCGC | TTGCACCAAG | CAGTGGGATG | TGGTAAGCTA | CCTCCTGGAG | AACCCACACC | AGCCCCCCAG | CCTGCAGGCC |
| flhX21ellrha2   | #50  | CTACCCCTCT | CTTTGGCCGC | TTGCACCAAG | CAGTGGGATG | TGGTAAGCTA | CCTCCTGGAG | AACCCACACC | AGCCCCCCAG | CCTGCAGGCC |
| flhX21ellrmv2   | #48  | CTACCCCTCT | CTTTGGCCGC | TTGCACCAAG | CAGTGGGATG | TGGTAAGCTA | CCTCCTGGAG | AACCCACACC | AGCCCCCCAG | CCTGCAGGCC |
| flhX21ellrlv1   | #47  | CTACCCCTCT | CTTTGGCCGC | TTGCACCAAG | CAGTGGGATG | TGGTAAGCTA | CCTCCTGGAG | AACCCACACC | AGCCCCCCAG | CCTGCAGGCC |
| frhoc12c4h3..   | #43  | CTACCCCTCT | CTTTGGCCGC | TTGCACCAAG | CAGTGGGATG | TGGTAAGCTA | CCTCCTGGAG | AACCCACACC | AGCCCCCCAG | CCTGCAGGCC |
| flhX21ellrlv2   | #40  | CTACCCCTCT | CTTTGGCCGC | TTGCACCAAG | CAGTGGGATG | TGGTAAGCTA | CCTCCTGGAG | AACCCACACC | AGCCCCCCAG | CCTGCAGGCC |
| flhX21ellrmv1   | #36  | CTACCCCTCT | CTTTGGCCGC | TTGCACCAAG | CAGTGGGATG | TGGTAAGCTA | CCTCCTGGAG | AACCCACACC | AGCCCCCCAG | CCTGCAGGCC |
| flhX21ellrja2   | #26  | CTACCCCTCT | CTTTGGCCGC | TTGCACCAAG | CAGTGGGATG | TGGTAAGCTA | CCTCCTGGAG | AACCCACACC | AGCCCCCCAG | CCTGCAGGCC |
| Gembank AA992.. | #9   | CTACCCCTCT | CTTTGGCCGC | TTGCACCAAG | CAGTGGGATG | TGGTAAGCTA | CCTCCTGGAG | AACCCACACC | AGCCCCCCAG | CCTGCAGGCC |
| Gembank AA995.. | #9   | CTACCCCTCT | CTTTGGCCGC | TTGCACCAAG | CAGTGGGATG | TGGTAAGCTA | CCTCCTGGAG | AACCCACACC | AGCCCCCCAG | CCTGCAGGCC |
| flhX21ellrr9..  | >#1> | CCCTCT     | CTTTGGCCGC | TTGCACCAAG | CAGTGGGATG | TGGTAAGCTA | CCTCCTGGAG | AACCCACACC | AGCCCCCCAG | CCTGCAGGCC |
| flhX21ellrr9..  | >#1> | CCCTCT     | CTTTGGCCGC | TTGCACCAAG | CAGTGGGATG | TGGTAAGCTA | CCTCCTGGAG | AACCCACACC | AGCCCCCCAG | CCTGCAGGCC |
| jthb051a01t..   | >#1> |            | CTTTGGCCGC | TTGCNCCAAG | CAGNGGGATG | TGGTAAGCTA | CCTNCTGGAG | AACCCACACC | AGNCCGCCAG | CCTGCAGGCC |
| flhX21ellrr1..  | >#1> |            |            |            |            | TA         | CCTCCTGGAG | AACCCACACC | AGCCCCCCAG | CCTGCAGGCC |
| flhX21ellrr1..  | >#1> |            |            |            |            | TA         | CCTCCTGGAG | AACCCACACC | AGCCCCCCAG | CCTGCAGGCC |

#991 CTACCCCTCT CTTGGCCGC TTGCACCAAG CAGTGGGATG TGTAAGCTA CCTCCTGGAG AACCCACACC AGCCCGCCAG CCTGCAGGCC  
L P L S L A A C T K Q W D V V S Y L L E N P H Q P A S L Q A  
+ + + + + + + + + + + + + + +

[illegible]

#1081 ACTGACTCCC AGGGCAACAC AGTCTGTCAT GCCCTAGTGA TGATCTCGGA CAACTCAGCT GAGAACATTG CACTGGTGAC CAGCATGTAT  
T D S Q G N T V L H A L V M I S D N S A E N I A L V T S M Y  
+ + + + +

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frhob12c4g1... #231 GATGGGCTCC TCCAAGCTGG GGGCCGCCTC TGCCTTACC GTCAGCTTGA GGACATCCGC AACCTNCAGG ATCTCACGCC TCTGAAGCTG
frhob012c04s... #231 GATGGGCTCC TCCAAGCTGG GGGCCGCCTC TGCCTTACC GTCAGCTTGA GGACATCCGC AACCTGCAGG ATCTCACGCC TCTGAAGCTG
frhob012c04s... #231 GATGGGCTCC TCCAAGCTGG GGGCCGCCTC TGCCTTACC GTCAGCTTGA GGACATCCGC AACCTGCAGG ATCTCACGCC TCTGAAGCTG
frhob12c4h3... #223 GATGGGCTCC TCCAAGCTGG GGGCCGCCTC TGCCTTACC GTCAGCTTGA GGACATCCGC AACCTGCAGG ATCTCACGCC TCTGAAGCTG
Genbank AA9... #189 GATGGGCTCC TCCAAGCTGG GGGCCGCCTC TGCCTTACC GTCAGCTTGA GGACATCCGC AACCTGCAGG ATCTCACG
Genbank AA9... #189 GATGGGCTCC TCCAAGCTGG GGGCCGCCTC TGCCTTACC GTCAGCTTGA GGACATCCGC AACCTGCAGG ATCTCACGCC TCTGAAGCTG
jthLa029c10t... >#1> CGCCTC TGCCTTACC GTCAGCTTGA GGACATCCGC AACCTGCAGG ATCTCACGCC TCTGAAGCTG
Genbank W38... >#1> CCTC TGCCTTACC GTCAGCTTGA GGACATCCGC AACCTGCAGG ATCTCACGCC TCTGAAANTG

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#1171 GATGGGCTCC TCCAAGCTGG GGGCCGCCTC TGCCTTACC GTCAGCTTGA GGACATCCGC AACCTGCAGG ATCTCACGCC TCTGAAGCTG
      D G L L Q A G A R L C P T V Q L E D I R N L Q D L T P L K L
                                     +
                                     ++

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frhob12c4g1... #321 GCCGCCAAGG AGGGCAAGNT CGANATTTTC AGGCACATCC TGCA
frhob012c04s... #321 GCCGCCAAGG AGGGCAAGAT CGAGATTTTC AGGCACATCC TGCAGCGGGA GTTTTCAGGA CTGAGCCACC TTTCGCCAAA GTTCACCGAG
frhob012c04s... #321 GCCGCCAAGG AGGGCAAGAT CGAGATTTTC AGGCACATCC TGCAGCGGGA GTTTTCAGGA CTGAGCCACC TTTCGCCAAA GTTCACCGAG
frhob12c4h3... #313 GCCGCCAAGG AGGGCAAGAT CGAGATTTTC AGGCACATCC TGCAGCGGGA GTTTTCAGGA CTGAGCCACC TTTCGCCAAA GTTCACCGAG
Genbank AA9... #279 GCCGCCAAGG AGGGCAAGAT CGAGATCTTC AGGCACATCC T
jthLa029c10t... #67 GCCGCCAAGG AGGGCAAGAT CGAGATTTTC AGGCACATCC TGCAGCGGGA GTTTTCAGGA CTGAGCCACC TTTCGCCAAA GTTCACCGAG
Genbank W386... #65 GCCGCCAAGG AGGGCAAGAT CGAGATTTTC AGGCACATCC TGCAGCGGGA GTTTTCAGGA CTGAGCCACC TTTCGCCAAA GTTCACCGAG
Genbank AA3... >#1> TTCAGGA CTGAGCCACC TTTCGCCAAA GTTCACCGAG

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#1261 GCCGCCAAGG AGGGCAAGAT CGAGATTTTC AGGCACATCC TGCAGCGGGA GTTTTCAGGA CTGAGCCACC TTTCGCCAAA GTTCACCGAG
      A A K E G K I E I F R H I L Q R E F S G L S H L S R K F T E
                + + +

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frhob012c04s... #411 TGGTGCTATG GGCCTGTCCG GGTGTGCTG TATGACCTGG CTTCTGTGGA CAGCTGTGAG GAGAACTCAG TGCTGGAGAT CATTGCCTTT
frhob012c04s... #411 TGGTGCTATG GGCCTGTCCG GGTGTGCTG TATGACCT:G CTTCTGTGGA CAGCTGTGAG GAGAACTCAA TGCTGGAGAT CATTGCCTTT
frhob12c4h3... #403 TGGTGCTATG GGCCTGTCCG GGTGTGCTG TATGACCTGG CTTCTGTGGA CAGCTGTGAG GAGAACTCAG TGCTGGAGAT CATTGCCTTT
jthLa029c10t... #157 TGGTGCTATG GGCCTGTCCG GGTGTGCTG TATGACCTGG CTTCTGTGGA CAGCTGTGAG GAGAACTCAG TGCTGGAGAT CATTGCCTTT
Genbank W38... #155 TGGTGCTATG GGCCTGTCCG GGTGTGCTG TATGACCTGG CTTCTGTGGA CAGCTGTGAG GAGAACTCAG TGCTGGAGAT CATTGCCTTT
Genbank AA35... #38 TGGTGCTATG GGCCTGTCCG GGTGTGCTG TATGACCTGG CTTCTNTGGA CAGCTGTNAG GAGAACTCAG TGCTGGAGAT CATTGCCTTT
frhob12c4c1... >#1> GGCCTGTNCG GGTGTNCTG TATGACNTGG CTTCTGTGNA CAGCTGTGAW GAGAACTCAG TGCTGGAGAT CATTGCCTTT
Genbank N24... >#1> GAGAACTCAG TGCTGGAGAT CATTGNCCTT

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#1351 TGGTGCTATG GGCCTGTCCG GGTGTGCTG TATGACCTGG CTTCTGTGGA CAGCTGTGAG GAGAACTCAG TGCTGGAGAT CATTGCCTTT
      W C Y G P V R V S L Y D L A S V D S C E E N S V L E I I A F
                + + + + +

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frhob012c04s... #501 CATTGCAAGA GCCCGCACCG ACACCGAATG GTCGNTTTGG AGCCCTGAA CAAACTGCTG C
frhob012c04s... #501 CATTGCAAGA ACCCGCACCG ACACCGAATG GTCGNTTTGG AGCCCTGAA CAAACTGCTG CANGCGAAAT GGGATCTGCT CAT:CCCCAAG
frhob12c4h3... #493 CATTGCAAGA GCCCGCACCG ACACCGAATG GTCGNTTTGG AGCCCTGAA CAAACTGCTG CANGCGAAAT GGGATCTGCT CATCCCCAA
jthLa029c10t... #247 CATTGCAAGA GCCCGNACCG ACACCGAATG GTCGNTTTGG AGCCCTGAA CAAACTGCTG CANGCGAAAT GGGATCTGCT CATNCCNCAAG
Genbank W38... #245 CATTGCAAGA GCCCGCACCG ACACCGAATG GTCGNTTTGG AGCCCTGAA CAAACTGCTG CANGCGAAAT GGGATCTGCT CATCCCCAAG
Genbank AA3... #128 CATTGCAAGA GCCCGCACCG ACACCGAATG GTCGNTTTGG AGCCCTGAA CAAACTGCTG CANGCGAAAT GGGATCTGCT CATCCCCAAG
frhob12c4c1... #81 CATTGCAAGA GCCCGCACCG ACACCGAATG GTCGNTTTGG AGCCCTGAA CAAACTGCTG CANGCGAAAT GGGATCTGCT CATCCCCAAG
Genbank N242... #31 CATTGCAAGA GCCCGNACCG ACACCGAATG GTCGNTTTGG AGCCCTGAA CAAACTGCTG CANGCGAAAT GGGATCTGCT CATCCCCAAG

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#1441 CATTGCAAGA GCCCGCACCG ACACCGAATG GTCGNTTTGG AGCCCTGAA CAAACTGCTG CANGCGAAAT GGGATCTGCT CATCCCCAAG
      H C K S P H R H R M V V L E P L N K L L Q A K W D L L I P K
                ++ + + +

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21e11a  
Sequencher™ "21e11racefinal"

Achr21e11 ... #425 GGCACCTAAGC CAGATGGCAG CCCGGATGAG CGCTGGTGCT TCAGGGTGA GGAGGTGAAC TGGGCTTCAT GGGAGCAGAC GCTGCCTACG  
frhob12c4d1... >#1> TCAT GGGAGCAGAC GCTGCCTACG

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#2431 GGCACCTAAGC CAGATGGCAG CCCGGATGAG CGCTGGTGCT TCAGGGTGA GGAGGTGAAC TGGGCTTCAT GGGAGCAGAC GCTGCCTACG  
G T K P D G S P D E R W C F R V E E V N W A S W E Q T L P T

---

Achr21e11 ... #515 CTGTGTGAGG ACCCGTCAGG GGCAGGTGTC CCTCGAACTC TCGAGAACCC TGTCTGGCT TCCCCTCCCA AGGAGGATGA GGATGGTGCC  
frhob12c4d1... #25 CTGTGTGAGG ACCCGTCAGG GGCAGGTGTC CCTCGAACTC TCGAGAACCC TGTCTGGCT TCCCCTCCCA AGGAGGATGA GGATGGTGCC

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#2521 CTGTGTGAGG ACCCGTCAGG GGCAGGTGTC CCTCGAACTC TCGAGAACCC TGTCTGGCT TCCCCTCCCA AGGAGGATGA GGATGGTGCC  
L C E D P S G A G V P R T L E N P V L A S P P K E D E D G A

+

Achr21e11 ... #605 TCTGAGGAAA ACTATGTGCC CGTCCAGCTC CTCCAGTCCA ACTGATGGCC CAGATGCAGC AGGAGGCCAG AGGACAGAGC AGAGGATCTT  
frhob12c4d1... #115 TCTGAGGAAA ACTATGTGCC CGTCCAGCTC CTCCAGTCCA ACTGATGGCC CAGATGCAGC AGGAGGCCAG AGGACAGAGC ANAGGATCTT

---

#2611 TCTGAGGAAA ACTATGTGCC CGTCCAGCTC CTCCAGTCCA ACTGATGGCC CAGATGCAGC AGGAGGCCAG AGGACAGAGC AGAGGATCTT  
S E E N Y V P V Q L L Q S N . W P R C S R R P E D R A E D L

+

Achr21e11 ... #695 TCCAACCACA TCTGCTGGCT CTGGGGTCCC AGTGAATTCT GGTGGCAAAT ATATATTTTC ACTAACTCAA AAAAAAAAAA AAAAA  
frhob12c4d1... #205 TCCAACCACA TCTGCTGGCT CTGGGGTCCC AGTGAATTCT GGTGGCAAAT ATATATTTTC ACTAACTCAA AAAAAAAAAA AAAAAAAAAA

---

#2701 TCCAACCACA TCTGCTGGCT CTGGGGTCCC AGTGAATTCT GGTGGCAAAT ATATATTTTC ACTAACTCAA AAAAAAAAAA AAAAAAAAAA  
S N H I C W L W G P S E F W W Q I Y I P T N S K K K K K K K

---

frhob12c4d1... #295 AAAAAAAAAA AAAAAAAAAA

---

#2791 AAAAAAAAAA AAAAAAAAAA  
K K K K K K

# Exhibit C

BLASTP vs. PNU (AA) flh2lellorfaa - 5:26:57 pm on Dec 21 98

BLASTP 1.4.10MP-WashU [30-Aug-96] (Build 20:24:58 Oct 21 1996)

Reference: Altschul, Stephen F., Warren Gish, Webb Miller, Eugene W. Myers, and David J. Lipman (1990). Basic local alignment search tool. J. Mol. Biol. 215:403-10.

Query= flh2lellorfaa  
(764 letters)

Database: /disk0/asap/databases/db/NRP/protxnu  
345,099 sequences; 106,000,184 total letters.

Searching.....done

| Sequences producing High-scoring Segment Pairs:              | High<br>Score | Smallest<br>Sum<br>Probability |   |
|--|---------------|--------------------------------|---|
|  |               | P(N)                           | N |
| GP:gi 2570933 (AF029310) vanilloid receptor subtype 1 [Ra... | 336           | 1.9e-221                       | 7 |
| GPU:gi 3986159 gnl PID d1035925 (AB015231) VR1sk [Mus mus... | 299           | 9.8e-136                       | 6 |
| GP:gi 2911863 (AF047660) contains similarity to ankyrin r... | 103           | 6.5e-20                        | 8 |
| GP:gi 3875319 gnl PID e1344970 (Z74030) similar to ankyri... | 97            | 1.1e-19                        | 7 |
| GP:gi 2642590 (AF031408) olfactory channel [Caenorhabditi... | 93            | 3.6e-16                        | 6 |
| GP:gi 2854148 (AF045639) contains similarity to ankyrin r... | 93            | 4.1e-16                        | 6 |
| GP:gi 3879753 gnl PID e1349345 (Z72514) Similarity to Hum... | 73            | 3.6e-11                        | 6 |
| GP:gi 3287188 gnl PID e315126 (Y10601) ankyrin-like prote... | 71            | 2.7e-08                        | 6 |
| SP:SP:sp P48994 TRPL_DROME TRANSIENT-RECEPTOR-POTENTIAL L... | 69            | 2.2e-05                        | 4 |
| GP:gi 1841966 (U65916) ankyrin [Rattus norvegicus]           | 72            | 5.6e-05                        | 3 |

WARNING: Descriptions of 4 database sequences were not reported due to the limiting value of parameter V = 10.

>GP:gi|2570933 (AF029310) vanilloid receptor subtype 1 [Rattus norvegicus]  
Length = 838

Score = 336 (156.0 bits), Expect = 1.9e-221, Sum P(7) = 1.9e-221  
Identities = 67/112 (59%), Positives = 84/112 (75%)

Query: 206 FYFGELPLSLAACTKQWDVVSXLLNPHQPASLQATDSQGNTVLHALVMISDNLSAENIAL 265  
FYFGELPLSLAACT Q +V +LL+N QPA + A DS GNTVLHALV ++DN+ +N  
Sbjct: 245 FYFGELPLSLAACTNQLAIVKFLQNSWQPADISARDSVGNLTVLHALVEVADNTVDNTRKF 304

Query: 266 VTSMYDGLLQAGARLCPTVQLEDIRNLQDLTPLKLAKEGKIEIFRHILORE 317  
VTSMY+ +L GA+L PT++LE+I N + LTPL LAA GKI + +ILORE  
Sbjct: 305 VTSMYNEILILGAKLHPTLKLKEITNRKGLTPLALAASSGKIGVLAYILORE 356

Score = 316 (146.7 bits), Expect = 1.9e-221, Sum P(7) = 1.9e-221  
Identities = 65/138 (47%), Positives = 93/138 (67%)

Query: 66 ASQPDPNRFRDRRLFNAVSRGVPEDLAGLPEYLSKTSKYLTDSEYTEGSTGKTCIMKAVL 125  
A + P +DR +F+AV++ ++L L +L ++ K LTDSE+ + TGKTCI+KA+L  
Sbjct: 104 AGEKFPRLYDRSIFDAVAQSNQCELESLLPFLQRSKKRLTDSEFKDPETGKTCILKAML 163

Query: 126 NLKDGVNACILPLLQIDRDSGNFQPLVNAQCTDDYYRGHSALHIAIEKRSLOQVLLLVEN 185  
NL +G N I LL + R + + + VNA TD YY+G +ALHIAIE+R++ V LLVEN  
Sbjct: 164 NLHNGQNDTIALLLDVARKTDSLKQFVNASVTDSSYYKGQTALHIAIERRNMTETVLLLVEN 223

Query: 186 GANVHARACGRFFQKGQG 203

GA+V A A G FF+K +G  
Sbjct: 224 GADVQAAANGDFFKKTG 241

Score = 299 (138.8 bits), Expect = 1.9e-221, Sum P(7) = 1.9e-221  
Identities = 69/175 (39%), Positives = 94/175 (53%)

Query: 391 FFLNFLCNLIYMFIFTAVAYHQPTLKAAPHLKAEVGNMMLLTGHILILLGGIYLLVGQL 450  
F+ NF +YM IFTA AY++P LK VG+ +TG IL + +

Sbjct: 434 FYFNFFVYCLYMIIFTAAAYRPFVEGLPPYKLNKNTVGDYFRVTGEILSVXXXXXXXRG I 493

Query: 451 WYFWRRHVFIWISFIDSIFEILFLFQALLKPVSVQLCFLAIEWYLP LLVSALV LGWLNLL 510  
YF +R + F+DSY EILF Q+L VS VL F + Y+ +V +L +GW N+L

Sbjct: 494 QYFLQRRPSLKSLEFVDSYSEILFFVQSLEFLVSVVLYFSQRKEYVASMVFS LAMGWTNML 553

Query: 511 YYTRGFQHTGIYSVMIQVILRDLRLFLLIYLVFLPGFAVALVSLSQEAWRPEAP 565  
YYTRGFQ GIY+VMI+K+ILRDL R A+V+L ++ P

Sbjct: 554 YYTRGFQQMGIYAVMIEKMILRDL CRXXXXXXXXXXXXXTAVVT LIEDGKNNSLP 608

Score = 281 (130.5 bits), Expect = 1.9e-221, Sum P(7) = 1.9e-221  
Identities = 54/95 (56%), Positives = 70/95 (73%)

Query: 586 GAQYRGILEASLELFKFTIGMGELAFQEQLHFRGMVLLLLLAYVLLTYILLNMLIALMS 645  
G Y + LELFKFTIGMG+L F E F+ + ++LLLAYV+LTYILLNMLIALM

Sbjct: 624 GNSYNSLYSTCLELFKFTIGMGDLEFTENYDFKAVFIILLLAYVILTYILLNMLIALMG 683

Query: 646 ETVNSVATDSWSIWKLQKAISVLEMENGYWWCRKK 680

ETVN +A +S +IWKLQ+AI++L+ E + C +K

Sbjct: 684 ETVNKIAQESKNIWKLQRAITILDTEKSFLKCMRK 718

Score = 162 (75.2 bits), Expect = 1.9e-221, Sum P(7) = 1.9e-221  
Identities = 28/38 (73%), Positives = 34/38 (89%)

Query: 323 HLSRKFTWCYGPVRVSLYDLASVDSCEENSVLEIIAF 360

HLSRKFTEW YGPV SLYDL+ +D+CE+NSVLE+IA+

Sbjct: 364 HLSRKFTEWAYGPFVHSSLYDLSCIDTCEKNSVLEVIAY 401

Score = 144 (66.9 bits), Expect = 1.9e-221, Sum P(7) = 1.9e-221  
Identities = 25/47 (53%), Positives = 31/47 (65%)

Query: 678 RKKQRAGVMLTVGTPDGSPDERWCFRVBEVNWASWEQTLPTLCEDP 724

RK R+G +L VG PDG D RWCFRV+EVNW +W + + EDP

Sbjct: 717 RKAFRSGKLLQVGFTPDGKDDYRWCFRVDEVNWTWNTNNGIINEDP 763

Score = 94 (43.6 bits), Expect = 1.9e-221, Sum P(7) = 1.9e-221  
Identities = 15/30 (50%), Positives = 24/30 (80%)

Query: 363 KSPHRHRMVLEPLNKLQAKWDLIPKFF 392

++P+RH M+++EPLN+LLQ KWD + + F

Sbjct: 405 ETPNRHDMLLVEPLNRLQDKWDRFVKRIF 434

Score = 46 (21.4 bits), Expect = 1.3e-162, Sum P(6) = 1.3e-162  
Identities = 11/48 (22%), Positives = 24/43 (50%)

Query: 435 HILILLGGIYLLVGQLWYFWRRHVFIWISFIDSIFEILFLFQALLKPV 482

H ++L+ + L+ W + + +F+ F+ + I+F A +PV

Sbjct: 410 HDMLLVEPLNRLQLQDKWDRFVKRIFYFNFFVYCLYMIIFTAAAYRPV 457

Score = 37 (17.2 bits), Expect = 2.4e-169, Sum P(7) = 2.4e-169  
Identities = 8/22 (36%), Positives = 13/22 (59%)

Query: 297 PLKLAKEGKIEIFRHILQREF 318

PL LAA ++ I + +LQ +

Sbjct: 251 PLSLAACNTQLAIVKFLQNSW 272

Score = 37 (17.2 bits), Expect = 1.9e-111, Sum P(7) = 1.9e-111  
Identities = 8/26 (30%), Positives = 15/26 (57%)

Query: 206 FYFGELPLSLAACTKQWDVVSYLEN 231  
+Y G+ L +A + +V+ L+EN  
Sbjct: 198 YYKGQTALHIAIERRNMTLVTLLEN 223

>GPU:gi|3986159|gnl|PID|d1035925 (AB015231) Vrlsk [Mus musculus]  
Length = 563

Score = 299 (138.8 bits), Expect = 9.8e-136, Sum P(6) = 9.8e-136  
Identities = 69/175 (39%), Positives = 94/175 (53%)

Query: 391 FFLNFCNLIIYMFIFTAVAYHQPTLKKAAAPHLKAEVGNMMLTGHILILLGGIYLLVGQL 450  
F+ NF +YM IFTA AY++P LK VG+ +TG IL + +  
Sbjct: 127 FYNFFVYCLYMIIFTAAAYRPVEGLPPYKLKNTVGDYFRVTGEILSVXXXXXXXXXRG 186

Query: 451 WYFWRRHVFIWISFIDSYPEILFLFQALPKVSVQVLCFLAIEWYLP LLVSALVLGWLNNL 510  
YF +R + F+DSY EILF Q+L VS VL F + Y+ +V +L +GW N+L  
Sbjct: 187 QYFLQRRPSLKSFLVDSYSEILFFVQSLFMLVSVVLYFSQRKEYVASMVFLAMGWTNML 246

Query: 511 YYTRGFQHTGIYSVMIQKVILRDLLRFLLIYLVFLFGFAVALVSLSQEAWRPEAP 565  
YYTRGFQ GIY+VMI+K+ILRD L A+V+L ++ P  
Sbjct: 247 YYTRGFQMGIIYAVMIEKMILRDLCRXXXXXXXXXXXXXTAVVTIEDGKNNSLP 301

Score = 253 (117.5 bits), Expect = 9.8e-136, Sum P(6) = 9.8e-136  
Identities = 50/89 (56%), Positives = 64/89 (71%)

Query: 586 GAQYRGILEASLELFKFTIGMGELAQEQHFRGMVLLLLLAYVLLTYILLNMLIALMS 645  
G Y + LELFKFTIGM+L F E F+ + ++LLLAYV+LTYILLNMLIALM  
Sbjct: 317 GNSYNSLYSTCLELFKFTIGMGDLFTENYDFKAVFIILLLAYVILTYILLNMLIALMG 376

Query: 646 ETVNSVATDSWSIWKLQKAISVLEMENGY 674  
ETV V+ +S IWKLQ A ++L++E +  
Sbjct: 377 ETVGQVSKESKHIWKLQWATTILDIERSF 405

Score = 163 (75.7 bits), Expect = 9.8e-136, Sum P(6) = 9.8e-136  
Identities = 28/47 (59%), Positives = 34/47 (72%)

Query: 678 RKKQRAGVMLTVGTPDGSPDERWCFRVEEVNWSWEQTLPTLCEDP 724  
RK R+G M+TVG DG+PD RWCFRV+EVNW+ W Q L + EDP  
Sbjct: 410 RKAFRSGEMTVGKSSDGTDDRWCFRVDEVNWSHWNQNLGIINEDP 456

Score = 162 (75.2 bits), Expect = 9.8e-136, Sum P(6) = 9.8e-136  
Identities = 28/38 (73%), Positives = 34/38 (89%)

Query: 323 HLSRKFTWCYGPVRVSLYDLASVDSCEENSVLEIIAF 360  
HLSRKFTEW YGPV SLYDL+ +D+CE+NSVLE+IA+  
Sbjct: 57 HLSRKFTEWAYGPVHSSLYDLSCIDTCEKNSVLEVIAY 94

Score = 129 (59.9 bits), Expect = 9.8e-136, Sum P(6) = 9.8e-136  
Identities = 27/49 (55%), Positives = 36/49 (73%)

Query: 269 MYDGLLQAGARLCPTVQLEDIRNLQDLTPLKLAKEGKIEIFRHILQRE 317  
MY+ +L GA+L PT++LE+I N + LTP LAA GKI + +ILQRE  
Sbjct: 1 MYNEILILGAKLHPTLKLEETNRKGLTPLALAASSGKIGVLAYILQRE 49

Score = 94 (43.6 bits), Expect = 9.8e-136, Sum P(6) = 9.8e-136  
Identities = 15/30 (50%), Positives = 24/30 (80%)

Query: 363 KSPHRHRMVVLEPLNKLQAKWDLIPKFF 392

++P+RH M+++EPLN+LLQ KWD + + F  
Sbjct: 98 ETPNRRHDMLLVEPLNRLQLQDKWDRFVKRI 127

Score = 46 (21.4 bits), Expect = 2.3e-77, Sum P(5) = 2.3e-77  
Identities = 11/48 (22%), Positives = 24/48 (50%)

Query: 435 HILILLGGIYLLVGQLWYFWRRHVFIWISFIDSYFEILFLFQALLKPV 482  
H ++L+ + L+ W + + +F + F+ + I+F A +PV  
Sbjct: 103 HDMLLVEPLNRLQLQDKWDRFVKRIFYFNFFVYCLYMIIFTAAAYRPV 150

Score = 42 (19.5 bits), Expect = 1.2e-122, Sum P(6) = 1.2e-122  
Identities = 9/22 (40%), Positives = 16/22 (72%)

Query: 209 GELPLSLAACTKQWDVVSYLLE 230  
G PL+LAA + + V++Y+L+  
Sbjct: 26 GLTPLALAASSGKIGVLAYILQ 47

>GP:gi|2911863 (AF047660) contains similarity to ankyrin repeats  
[Caenorhabditis elegans]  
Length = 900

Score = 103 (47.8 bits), Expect = 6.5e-20, Sum P(8) = 6.5e-20  
Identities = 23/89 (25%), Positives = 44/39 (49%)

Query: 497 LLVSALVLGWLNLXYTRGFQHTGIYSVMIQVILRDLLRFLLIYLVFLFGFAVALVSL 556  
L+ ++ ++ LYY R + G + +M+ +I D+ RF+LIY +FL GF+ + +  
Sbjct: 592 LITVTMIFTTVHYLYYCRVIRFVGPFVLMVYTIATDIFRMLIYGIFLMGFSQSFSLLIF 651

Query: 557 QEAWRPEAPTGPNATESVQPMEGQEDEGN 585  
R T+ + EG +++ N  
Sbjct: 652 LSCERBANVIKKLITDQSEASEGSDNKN 680

Score = 63 (29.3 bits), Expect = 6.5e-20, Sum P(8) = 6.5e-20  
Identities = 15/27 (55%), Positives = 18/27 (66%)

Query: 291 NLQDLTPLKLAKEGKIEIFRHILQRE 317  
N Q L+PL LAAK K E+F IL+ E  
Sbjct: 328 NKQSLSPFLTAAKLAKKEMFDEILELE 354

Score = 56 (26.0 bits), Expect = 6.5e-20, Sum P(8) = 6.5e-20  
Identities = 16/48 (33%), Positives = 25/48 (52%)

Query: 151 LVNAQCTDDYYRGHSALHIAIEKRSLOQCVLLVENGANVHARACGRFF 198  
L+N + + G S LH AI + V ++ GA+V++R G FF  
Sbjct: 185 LLNDIHISEDYGLSPLHQAIINTDCKLVYKFLKLGADVNSRCYGAF 232

Score = 54 (25.1 bits), Expect = 6.5e-20, Sum P(8) = 6.5e-20  
Identities = 12/23 (52%), Positives = 13/23 (56%)

Query: 207 YFGELPLSLAACTKQWDVVSYLL 229  
Y GE PLS AAC Q + LL  
Sbjct: 263 YLGEYPLSFAACLNQPEFRLL 285

Score = 49 (22.8 bits), Expect = 6.5e-20, Sum P(8) = 6.5e-20  
Identities = 8/23 (34%), Positives = 12/23 (52%)

Query: 331 WCYGPVRVSLYDLASVDSCEENS 353  
W YG + Y LA +D+ E +  
Sbjct: 359 WAYGDASSTAYPLAKIDTINETT 381

Score = 47 (21.8 bits), Expect = 6.5e-20, Sum P(8) = 6.5e-20  
Identities = 11/30 (36%), Positives = 18/30 (60%)

Query: 236 ASLQATDSQGNTVLHALVMISDNSENIAL 265  
A+ A D+ GN+VLH V+ + + +AL  
Sbjct: 289 ANPNAQDTNGNSVLHMCVIHENMAMFKLAL 318

Score = 46 (21.4 bits), Expect = 5.7e-13, Sum P(6) = 5.7e-13  
Identities = 9/27 (33%), Positives = 17/27 (62%)

Query: 163 GHSALHIAIEKRSLQCVKLLVENGANV 189  
G+S LH+ + ++ KL +E GA++  
Sbjct: 298 GNSVLHMCVIHENMAMFKLALECGASL 324

Score = 43 (20.0 bits), Expect = 6.5e-20, Sum P(8) = 6.5e-20  
Identities = 8/18 (44%), Positives = 13/18 (72%)

Query: 634 ILLLNMLIALMSETVNSV 651  
I+ NMLIA+M+ T ++  
Sbjct: 753 IMQFNMLIAMMTRTYETI 770

Score = 41 (19.0 bits), Expect = 6.5e-20, Sum P(8) = 6.5e-20  
Identities = 10/27 (37%), Positives = 16/27 (59%)

Query: 113 GSTGKTCLMKAVLNKDGVNACILPLL 139  
GS G+T + +L+ D NA +L +L  
Sbjct: 153 GSMGETIIGCCLLHASDIHNALVLKIL 179

Score = 35 (16.3 bits), Expect = 0.00016, Sum P(3) = 0.00016  
Identities = 9/37 (24%), Positives = 16/37 (43%)

Query: 67 SQPDPNRFDRDLFNAVSRGVPEDLAGLPEYLSKTSK 103  
S+P P R+ ++ V + GL E+ S+  
Sbjct: 464 SEFPFGRYGNSTLQQVKPVINATSRGLVEWSEPLSQ 500

Score = 34 (15.8 bits), Expect = 6.8e-11, Sum P(8) = 6.8e-11  
Identities = 8/31 (25%), Positives = 17/31 (54%)

Query: 422 LKAEVGNMMLLTGHILILLGGIYLLVGQLWY 452  
L ++ +L+ ++LI + I+ V L+Y  
Sbjct: 577 LACDLSPVLLVVDNVLITVTMIFTTVHYLYY 607

>GP:gi|3875319|gnl|PID|e1344970 (Z74030) similar to ankyrin repeats  
[Caenorhabditis elegans] >GP:gi|3876480|gnl|PID|e1346172 (Z72508)  
similar to ankyrin repeats [Caenorhabditis elegans]  
Length = 790

Score = 97 (45.0 bits), Expect = 1.1e-19, Sum P(7) = 1.1e-19  
Identities = 18/53 (33%), Positives = 32/53 (60%)

Query: 497 LLVSALVLGWLNLLYYTRGFQHTGIYSVMIQVILRDLLRFLLIYLVFLFGFA 549  
+ + +++L + LYY R G + +M+ +I DL+RF +IY +FL GF+  
Sbjct: 526 MAIISILLVTQHFLYYMRAIPFVGPFVLMVYTTIATDLVRFAMIYSIFLVGFS 578

Score = 89 (41.3 bits), Expect = 1.1e-19, Sum P(7) = 1.1e-19  
Identities = 19/48 (39%), Positives = 27/48 (56%)

Query: 151 LVNAQCTDDYYRGHSALHIAIEKRSLQCVKLLVENGANVHARACGRFF 198  
L+N C + Y G S LH+AI + Q LL+ GA+++ R G FF  
Sbjct: 189 LINDICVSEYYGLSPLHLAIVNQDAQFTSLRLRLGADLNQRCYGAFF 236

Score = 62 (28.8 bits), Expect = 1.1e-19, Sum P(7) = 1.1e-19  
Identities = 13/23 (56%), Positives = 14/23 (60%)



Query: 207 YFGELPLSLAACTKQWDVVSYLL 229  
YFGE PLS A C Q D+ LL  
Sbjct: 267 YFGEYPLSFAICMGQHDLFRMLL 289

Score = 51 (23.7 bits), Expect = 1.1e-19, Sum P(7) = 1.1e-19  
Identities = 10/19 (52%), Positives = 13/19 (68%)

Query: 236 ASLQATDSQGNLVLHALVM 254  
A+L A D+ GNT LH V+  
Sbjct: 293 ANLSAQDTNGNTALHLCVI 311

Score = 49 (22.8 bits), Expect = 1.1e-19, Sum P(7) = 1.1e-19  
Identities = 11/51 (21%), Positives = 27/51 (52%)

Query: 601 KFTIGMGELAFQEQHLFRGMVLLLLLAYVLLTYILLNMLIALMSETVNSV 651  
+F++ E++ + + + L+ + + IL N+LIA+M+ T ++  
Sbjct: 626 EFSVLYREMSACDNFWMKWIGKLIFVIFETPVSILOFNLLIAMMTRTYETI 676

Score = 43 (20.0 bits), Expect = 1.2e-06, Sum P(5) = 1.2e-06  
Identities = 7/27 (25%), Positives = 16/27 (59%)

Query: 163 GHSALHIAIEKRSLQCVKLLVENGANV 189  
G++ALH+ + + + ++E G N+  
Sbjct: 302 GNTALHLCVHDKMDMLDAVLEAGGNI 328

Score = 39 (18.1 bits), Expect = 5.5e-12, Sum P(7) = 5.5e-12  
Identities = 12/56 (21%), Positives = 25/56 (44%)

Query: 461 WISFIDSYPEILFLFQALLKPVSQVLCFLAIEWYLP LLVSALVLGWLNL LYYTRGF 516  
W +F+ ++ L A L + + C LA ++ L + +++L T+ F  
Sbjct: 483 WFNFLKAFPAKLMFKGAFLEFIIISIPCLACSFHEFFLTIDNTMAIISILLVTQHF 538

Score = 38 (17.6 bits), Expect = 1.1e-19, Sum P(7) = 1.1e-19  
Identities = 9/18 (50%), Positives = 12/18 (66%)

Query: 289 IRNLQDLTPLKLAKEGK 306  
+ N Q+LT L LAA+ K  
Sbjct: 330 LANKQNLTA TLARLAK 347

Score = 37 (17.2 bits), Expect = 4.7e-18, Sum P(7) = 4.7e-18  
Identities = 7/15 (46%), Positives = 9/15 (60%)

Query: 723 DPSGAGVPRTLENPV 737  
DP G+ +ENPV  
Sbjct: 599 DPMGSEFNNIMENPV 613

Score = 36 (16.7 bits), Expect = 1.1e-19, Sum P(7) = 1.1e-19  
Identities = 10/47 (21%), Positives = 16/47 (34%)

Query: 376 LNKLLQAKWDL LIPKFFLNFLCNLIYMFIFTAVAYHQPTLKKAAAPHL 422  
+ ++L KW +L L I+ + AY HL  
Sbjct: 362 IEQILDEKWKAYGRALWLRSL L GFIFFYCCFVCAYMLRPSSATTEHL 408

Score = 35 (16.3 bits), Expect = 3.0e-08, Sum P(6) = 3.0e-08  
Identities = 9/23 (39%), Positives = 11/23 (47%)

Query: 207 YFGELPLSLAACTKQWDVVSYLL 229  
Y+G PL LA + S LL  
Sbjct: 199 YYGLSPLHLAIVNQDAQFTSLLL 221

Score = 34 (15.8 bits), Expect = 3.9e-12, Sum P(5) = 3.9e-12  
Identities = 7/21 (33%), Positives = 13/21 (61%)

Query: 297 PLKLAKEGKIEIFRHILORE 317  
PL A G+ ++FR +L ++  
Sbjct: 272 PLSFAICMGQHDLFRMLLAKK 292

Score = 34 (15.8 bits), Expect = 8.9e-10, Sum P(7) = 8.9e-10  
Identities = 6/18 (33%), Positives = 12/18 (66%)

Query: 610 AFQEQLHFRGMVLLLLLLA 627  
AF +L F+G L ++++  
Sbjct: 489 AFPKLMFKGAFLFIIIS 506

Score = 34 (15.8 bits), Expect = 4.0e-08, Sum P(6) = 4.0e-08  
Identities = 11/40 (27%), Positives = 16/40 (40%)

Query: 153 NAQCTDDYYRGHSALHIAIEKRSLOQCVKLLVENGANVHAR 192  
N T Y G L AI ++L+ AN+ A+  
Sbjct: 259 NTNYTGSMYFGEYPLSFAICMGQHDLFRMLLAKKANLSAQ 298

>GP:gi|2642590 (AF031408) olfactory channel [Caenorhabditis elegans]  
Length = 937

Score = 93 (43.2 bits), Expect = 6.8e-16, Sum P(5) = 6.8e-16  
Identities = 19/36 (52%), Positives = 25/36 (69%)

Query: 163 GHSALHIAIEKRSLOQCVKLLVENGANVHARACGRFF 198  
G SALH+AI + V LL+ + A+V+ARACG FF  
Sbjct: 172 GQSALHLAIVHDDYETVSLLLNSKADVNARACGNFF 207

Score = 92 (42.7 bits), Expect = 6.8e-16, Sum P(5) = 6.8e-16  
Identities = 20/53 (37%), Positives = 31/53 (58%)

Query: 497 LLVSALVLGWLNLNLLYYTRGFQHTGIYSVMIQKVILRDLLRFLLIYLVFLFGFA 549  
L V AL W+ LL++ R + TG + MI +I D++RF +I +FL F+  
Sbjct: 500 LEVFALPGSWIFLLFFARS AKLTGPFVQMIYSMIAGDMIRFAIISAIFLVSFS 552

Score = 57 (26.5 bits), Expect = 6.8e-16, Sum P(5) = 6.8e-16  
Identities = 10/24 (41%), Positives = 15/24 (62%)

Query: 207 YFGELPLSLAACTKQWDVVSYLLE 230  
Y+GE PL+ AAC D+ L++  
Sbjct: 226 YYGEYPLAFAACFGNKDIYDILLIQ 249

Score = 50 (23.2 bits), Expect = 6.8e-16, Sum P(5) = 6.8e-16  
Identities = 11/25 (44%), Positives = 15/25 (60%)

Query: 291 NLQDLTPLKLAKEGKIEIFRHILO 315  
N TPL LA K G+ +IF +L+  
Sbjct: 294 NHAGFTPLTLATKLGKQIFEEMLE 318

Score = 44 (20.4 bits), Expect = 6.8e-16, Sum P(5) = 6.8e-16  
Identities = 10/24 (41%), Positives = 15/24 (62%)

Query: 242 DSQGNLTVLHALVMISDNSAENIAL 265  
DS GNT+LH V+ +S + A+  
Sbjct: 258 DSFGNTILHMCVINYSSSMYSYAV 281

Score = 37 (17.2 bits), Expect = 2.2e-05, Sum P(4) = 2.2e-05  
Identities = 9/24 (37%), Positives = 13/24 (54%)

Query: 208 FGELPLSLAACTKQWDVVSYLLEN 231  
FG+ L LA ++ VS LL +  
Sbjct: 171 FGQSALHLAIVHDDYETVSLLLNS 194

Score = 34 (15.8 bits), Expect = 3.6e-16, Sum P(6) = 3.6e-16  
Identities = 7/21 (33%), Positives = 11/21 (52%)

Query: 364 SPHRHRMVVLEPLNKLQAKW 384  
+P M+ E + +LL KW  
Sbjct: 363 TPEHLDMIGSEVIQRLADKW 383

>GP:gi|2854148 (AF045639) contains similarity to ankyrin repeats  
[Caenorhabditis elegans]  
Length = 957

Score = 93 (43.2 bits), Expect = 7.6e-16, Sum P(5) = 7.6e-16  
Identities = 19/36 (52%), Positives = 25/36 (69%)

Query: 163 GHSALHTAIEKRSLOQCVKLLVENGANVHARACGRFF 198  
G SALH+AI + V LL+ + A+V+ARACG FF  
Sbjct: 175 GQSALHLAIVHDDYETVSLLLNSKADVNRACGNFF 210

Score = 92 (42.7 bits), Expect = 7.6e-16, Sum P(5) = 7.6e-16  
Identities = 20/53 (37%), Positives = 31/53 (58%)

Query: 497 LLVSALVLGWLNLLEYTRGFQHTGIYSVMIOKVILRDLLRFLLIYLVFLFGFA 549  
L V AL W+ LL++ R + TG + MI +I D++RF +I +FL F+  
Sbjct: 503 LFFVLPALPGSWIFLLFFARSALKTGPFVQMIYSMIAGDMIRFAIISAIPLVSFS 555

Score = 57 (26.5 bits), Expect = 7.6e-16, Sum P(5) = 7.6e-16  
Identities = 10/24 (41%), Positives = 15/24 (62%)

Query: 207 YFGELPLSLAACTKQWDVVSYLE 230  
Y+GE PL+ AAC D+ L++  
Sbjct: 229 YYGEYPLAFAACFGNKDIYDLILQ 252

Score = 50 (23.2 bits), Expect = 7.6e-16, Sum P(5) = 7.6e-16  
Identities = 11/25 (44%), Positives = 15/25 (60%)

Query: 291 NLQDLTPLKLAKEGKIEIFRHLQ 315  
N TPL LA K G+ +IF +L+  
Sbjct: 297 NHAGFTPLTLATKLGRKQIFPEMLE 321

Score = 44 (20.4 bits), Expect = 7.6e-16, Sum P(5) = 7.6e-16  
Identities = 10/24 (41%), Positives = 15/24 (62%)

Query: 242 DSQGNLVLHALVMISDNSENIAL 265  
DS GNT+LH V+ +S + A+  
Sbjct: 261 DSFGNTILHMCVINYSMYSYAV 284

Score = 37 (17.2 bits), Expect = 2.4e-05, Sum P(4) = 2.4e-05  
Identities = 9/24 (37%), Positives = 13/24 (54%)

Query: 208 FGELPLSLAACTKQWDVVSYLEN 231  
FG+ L LA ++ VS LL +  
Sbjct: 174 FGQSALHLAIVHDDYETVSLLLNS 197

Score = 34 (15.8 bits), Expect = 4.1e-16, Sum P(6) = 4.1e-16  
Identities = 7/21 (33%), Positives = 11/21 (52%)

Query: 364 SPHRHRMVVLEPLNKLQAKW 384  
+P M+ E + +LL KW  
Sbjct: 366 TPEHLDMIGSEVIQRLADKW 386

>GP:gi|3879753|gnl|PID|e1349345 (Z72514) Similarity to Human ankyrin  
(SW:ANK1\_HUMAN) [Caenorhabditis elegans]  
Length = 519

Score = 73 (33.9 bits), Expect = 3.6e-11, Sum P(6) = 3.6e-11  
Identities = 17/36 (47%), Positives = 19/36 (52%)

Query: 163 GHSALHIAIEKRSLQCVKLLVENGANVHARACGRFF 198  
G S L H A I L+ V L GA+VH R G FF  
Sbjct: 186 GLSPLHQAIVNEDLEMVYFLCRKGADVHQRCYGSFF 221

Score = 63 (29.3 bits), Expect = 3.6e-11, Sum P(6) = 3.6e-11  
Identities = 12/17 (70%), Positives = 13/17 (76%)

Query: 207 YFGELPLSLAACTKQWD 223  
Y+GE PLS AACT Q D  
Sbjct: 252 YWGEYPLSFAACTNQVD 268

Score = 52 (24.1 bits), Expect = 3.6e-11, Sum P(6) = 3.6e-11  
Identities = 13/27 (48%), Positives = 17/27 (62%)

Query: 289 IRNLQDLTPLKLAKEGKIEIFRHILQ 315  
+RN LTPL LAA+ K I+ IL+  
Sbjct: 315 VRNNLKLTPALAAARLAKKHIYDLILE 341

Score = 51 (23.7 bits), Expect = 3.6e-11, Sum P(6) = 3.6e-11  
Identities = 10/22 (45%), Positives = 12/22 (54%)

Query: 331 WCYGPVVRVSLYDLASVDSCEEN 352  
W YGPV Y L VD+ E+  
Sbjct: 348 WRYGPVVCKAYPLNDVDTINES 369

Score = 43 (20.0 bits), Expect = 3.6e-11, Sum P(6) = 3.6e-11  
Identities = 8/13 (61%), Positives = 10/13 (76%)

Query: 242 DSQGNTVLHALVM 254  
D+ GNTVLH V+  
Sbjct: 284 DTNGNTVLHLTVI 296

Score = 37 (17.2 bits), Expect = 3.6e-11, Sum P(6) = 3.6e-11  
Identities = 8/35 (22%), Positives = 16/35 (45%)

Query: 376 LNKLLQAKWDLIPKFFLNFLCNLIYMFIFQAVAY 410  
+ ++L++KW+ K L IY +A+  
Sbjct: 398 IEEVLESKWETFGKKQLFMSLAGYIYFLAVFYLA 432

>GP:gi|3287188|gnl|PID|e315126 (Y10601) ankyrin-like protein [Homo sapiens]  
Length = 1119

Score = 71 (33.0 bits), Expect = 2.7e-08, Sum P(6) = 2.7e-08  
Identities = 13/30 (43%), Positives = 20/30 (66%)

Query: 163 GHSALHIAIEKRSLQCVKLLVENGANVHAR 192  
G++ LH A+EK ++ VK L+ GAN + R  
Sbjct: 98 GNTPLHCAVEKNQIESVKFLLSRGANPNLR 127

Score = 63 (29.3 bits), Expect = 2.7e-08, Sum P(6) = 2.7e-08  
Identities = 10/30 (33%), Positives = 20/30 (66%)

Query: 501 ALVLGWLNLLEYTRGFQHTGIYSVMIQKVI 530  
A+ W+N L Y + F++ GI+ VM++ ++  
Sbjct: 838 AVYFYWMNFLLYLQRFENCIGIFIVMLEVIL 867

Score = 59 (27.4 bits), Expect = 9.0e-07, Sum P(6) = 9.0e-07  
Identities = 10/30 (33%), Positives = 19/30 (63%)

Query: 167 LHIAIEKRSIQCVKLLVENGANVHARACGR 196  
LH+A++ L+ +K+ ++NGA + GR  
Sbjct: 243 LHLAVQNGDLEMIKMCLDNGAQIDPVEKGR 272

Score = 50 (23.2 bits), Expect = 2.7e-08, Sum P(6) = 2.7e-08  
Identities = 10/26 (38%), Positives = 18/26 (69%)

Query: 291 NLQDLTFLKLAKEGKIEIFRHILQR 316  
+L +TPL LAAK G ++ + +L++  
Sbjct: 479 DLHGMPHLHLAAKNGHDKVQVQLLLKK 504

Score = 49 (22.8 bits), Expect = 2.7e-08, Sum P(6) = 2.7e-08  
Identities = 10/26 (38%), Positives = 14/26 (53%)

Query: 209 GELPLSLAACTKQWDVVSYLENPHQ 234  
G PL LA + W++V+ LL Q  
Sbjct: 342 GRSPLILATASASWNIVNLLLSKGAQ 367

Score = 48 (22.3 bits), Expect = 3.2e-05, Sum P(5) = 3.2e-05  
Identities = 9/32 (28%), Positives = 20/32 (62%)

Query: 524 VMIQKVILRDLLRFLLIYLVFLFGFAVALVSL 555  
+++ +VIL+ LLR ++++ L F ++ L  
Sbjct: 860 IVMLEVILKTLRSTVVFIPLLLAFGLSFYIL 891

Score = 41 (19.0 bits), Expect = 2.7e-08, Sum P(6) = 2.7e-08  
Identities = 9/28 (32%), Positives = 15/28 (53%)

Query: 525 MIQKVILRDLLRFLLIYLVFLFGFAVAL 552  
+I K +LR + F+ + L F F + L  
Sbjct: 865 VILKTLRSTVVFIPLLLAFGLSFYILL 892

Score = 40 (18.6 bits), Expect = 2.9e-07, Sum P(5) = 2.9e-07  
Identities = 13/50 (26%), Positives = 24/50 (48%)

Query: 620 MVLILLLLAYVLLTYILLNMLIALMSETVNSVATDSWSIWKLQKAISVLE 669  
+V + LL L++ +LLN+ S ++ + T S + + S LE  
Sbjct: 875 VVFIFILLAFGLSFYILLNLQDPFSSPLLSIIQTFSMMLGDINYRESFLE 924

Score = 40 (18.6 bits), Expect = 7.6e-05, Sum P(5) = 7.6e-05  
Identities = 8/19 (42%), Positives = 12/19 (63%)

Query: 296 TPLKLAKEGKIEIFRHIL 314  
TPL LA + G +E+ + L  
Sbjct: 241 TPLHLAVQNGDLEMIKMCL 259

Score = 39 (18.1 bits), Expect = 2.7e-08, Sum P(6) = 2.7e-08  
Identities = 7/19 (36%), Positives = 14/19 (73%)

Query: 625 LLAYVLLTYILLNMLIAL 643  
L+++ + I+L+N+LI L  
Sbjct: 941 LVSFTIFVPFIVLMNLLIGL 959

Score = 39 (18.1 bits), Expect = 6.8e-07, Sum P(6) = 6.8e-07  
Identities = 9/36 (25%), Positives = 18/36 (50%)

Query: 293 QDLTPLKLAKEGKIEIFRHILQREFSGLSHLSRK 328  
+ LT L + +IE+ H + +E+ + L+ F  
Sbjct: 681 EPLTALNAMVQNNRIELNHPVCKEYLLMKWLAYGF 716



## Qualified Target Summary Sheet

|  |   |
|--|---|
| <b>Target Name:</b>                    | Vanilloid Receptor Homologue (VR-2)   |
| <b>CFA Disease Area (s):</b>           | Pain  |
| <b>Druggable Target Class:</b>         | Calcium channel   |
| <b>MPI Gene Sequence Identifier:</b>   | Flh21e11 (Mine 18560)   |
| <b>Top Blast Hit:</b>                  | Rat vanilloid receptor 1 (VR1)  |
| <b>Amino Acid Coding Region:</b>       | ORF: 361-2652   |
| <b>CDNA Length:</b>                    | 2806 bp   |
| <b>Source (Tissue / Cell Line):</b>    | First clone identified in an internal heart library   |
| <b>Novel / Unrecognized:</b>           | ? Unrecognized / Novel  |
| <b>% Novelty:</b>                      | 66% novel across the complete cDNA.<br>Hits unannotated sequence in Non-Public Patent Data Base   |
| <b>Patent Status:</b>                  | Filed Nov. 1, 1998  |
| <b>Full Length Clone:</b>              | Yes   |
| <b>Expression Profiling Results:</b>   | Present in a sub-population of sensory neurons different from VR1.<br>Also present in sympathetic neurons.  |
| <b>Background:</b>                     | The published vanilloid receptor (Caterina et. Al. Nature 389:816-24, 1997) responds to heat and capsaicin by activating Ca++ influx in sensory neurons (Tominaga et.al., Neuron 21: 531-43, 1998). Capsaicin also binds to this channel. |
| <b>Hypothesis:</b>                     | This channel may be responsible for hypersensitivity in chronic neuropathic pain and represents a unique target for pain.   |
| <b>Assay Type:</b>                     | Cell-based assay  |
| <b>Reagents Needed:</b>                | Open reading frame will be cloned by Millennium into pCDNA 3.1  |
| <b>Readout:</b>                        |   |
| <b>Critical Experiments Necessary:</b> |   |

**MPI Target Name:**

Vanilloid Receptor Homologue (VR-2)

**QT Nomination Date:**

12/22/98

**Acti n Tak n:**

Accepted QT (unrecognized) Pending full length cDNA

**Date Accepted:**

**Bay r QT Leader:**

**MPI QT Leader:**

Rory Curtis or Peter DiStefano